

**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

In re Application of:	)	
	)	
<b>Robert DOBLHOFER et al.</b>	)	
	)	Group Art Unit: <b>Not Yet Assigned</b>
Application No.: <b>10/584,996</b>	)	
	)	Examiner: <b>Not Yet Assigned</b>
Filed: <b>June 29, 2006</b>	)	
	)	
For: <b>4-AMINO-7,8-DIHYDROPTERIDINES,</b>	)	
<b>PHARMACEUTICAL COMPOSITIONS</b>	)	
<b>CONTAINING THEM AND THEIR USE FOR</b>	)	
<b>THE TREATMENT OF DISEASES WHICH</b>	)	
<b>ARE CAUSED BY AN INCREASED NITRIC</b>	)	
<b>OXIDE LEVEL</b>	)	

**Commissioner for Patents**  
**P.O. Box 1450**  
**Alexandria, VA 22313-1450**

Sir:

**INFORMATION DISCLOSURE STATEMENT UNDER 37 C.F.R. § 1.97(b)**

Pursuant to 37 C.F.R. §§1.56 and 1.97(b), applicants bring to the Examiner's attention the documents listed on attached Form PTO/SB/08 and cited in the international search report. With exception of the U.S. patent, copies of the listed documents are attached. Applicants respectfully request that the Examiner consider the documents listed on attached Form PTO/SB/08 and indicate that they were considered by making an appropriate notation on this form. This Information Disclosure Statement is being filed before the mailing date of a first Office Action on the merits for the above-referenced application.

The following are listed on the accompanying PTO/SB/08 and are in a non-English language:

1. WO 95/32203 - An English language abstract of this document is enclosed.
2. DE 44 18 097
3. EP 0 906 913 A1
4. WO 01/21619 A1 - An English language abstract of this document is enclosed.
5. PFLEIDERER, W. et al., "A Simple Synthetic Approach to 8-Substituted 5,6,7,8-Tetrathyro- and 7,8-Dihydropterins," Chem. Ber., Vol. 104, pp. 2293-2312, (1971) - An English-language translation of the document is enclosed.
6. SCHIRCKS, V. B. et al., "A New, Regiospecific Synthesis of L-Biopterine," Helvetica Chimica Acta, Vol. 60, No. 1, pp. 211-214, (1977) - An English-language abstract of the document is enclosed..

In lieu of a statement of relevance or the translation of the non-English documents 2 and 3, enclosed is an English-language international search report from the Patent Office in the PCT international application, from which this national phase U.S. application is derived, citing these documents and setting forth the relevance thereof.

This submission does not represent that a search has been made or that no better art exists and does not constitute an admission that each or all of the listed documents are material or constitute "prior art." If the Examiner applies any of the documents as prior art against any claim in the application and applicants determine

that the cited documents do not constitute "prior art" under United States law, applicants reserve the right to present to the Office the relevant facts and law regarding the appropriate status of such documents. Applicants further reserve the right to take appropriate action to establish the patentability of the disclosed invention over the listed documents, should one or more of the documents be applied against the claims of the present application.

If there is any fee due in connection with the filing of this Statement, please charge the fee to our Deposit Account No. 06-0916.

Respectfully submitted,

FINNEGAN, HENDERSON, FARABOW,  
GARRETT & DUNNER, L.L.P.

Dated: June 11, 2007

By: 

Ernest F. Chapman  
Reg. No. 25,961

Enclosures  
EFC/FPD/sci

IDS Form PTO/SB/08: Substitute for form 1449A/PTO				<b>Complete if Known</b>	
<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  (Use as many sheets as necessary)				Application Number	10/584,996
				Filing Date	June 29, 2006
				First Named Inventor	Robert DOBLHOFFER et al.
				Art Unit	
				Examiner Name	
Sheet	1	of	2	Attorney Docket Number	05281.0018

U.S. PATENTS AND PUBLISHED U.S. PATENT APPLICATIONS					
Examiner Initials	Cite No. <sup>1</sup>	Document Number	Issue or Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number-Kind Code <sup>2</sup> (if known)			
		US-5,922,713	07-13-1999	WERNER	

**Note: Copies of the U.S. Patent Documents are not Required in IDS filed after October 21, 2004**

FOREIGN PATENT DOCUMENTS						
Examiner Initials	Cite No. <sup>1</sup>	Foreign Patent Document	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	Translation <sup>6</sup>
		Country Code <sup>3</sup> Number <sup>4</sup> Kind Code <sup>5</sup> (if known)				
		WO 95/32203	11-30-1995	PFLEIDERER et al.		Abstract
		DE 44 18 097 A1	11-30-1995	PFLEIDERER et al.		NO
		EP 0 906 913 A1	04-07-1999	WERNER et al.		NO
		WO 01/21619 A1	03-29-2001	PFLEIDERER et al.		Abstract
		WO 00/39129	07-06-2000	WAER et al.		
		GB 2 240 041 A1	07-24-1991	BRAQUET et al.		
		WO 93/13055	07-08-1993	BEAMS et al.		

NON PATENT LITERATURE DOCUMENTS			
Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	Translation <sup>6</sup>
		McCALL, T. B. et al., "Identification of N-Iminoethyl-L-Ornithine as an Irreversible Inhibitor of Nitric Oxide Synthase in Phagocytic Cells," Br. J. Pharmacol., Vol. 102, No. 1, p. 234, (1991). (Abstract Only)	
		MISKO, T. P. et al., "Selective Inhibition of the Inducible Nitric Oxide Synthase by Aminoguanidine," Eur. J. Pharmacol., Vol. 233, No. 1, p. 119, (1993). (Abstract Only)	
		MOORE, P. K. et al., "7-Nitro Indazole, an Inhibitor of Nitric Oxide Synthase, Exhibits Anti-Nociceptive Activity in the Mouse Without Increasing Blood Pressure," Br. J. Pharmacol., Vol. 108, No. 2, p. 296, (1993). (Abstract Only)	
		KWON, N. S. et al., "Reduced Biopterin as a Cofactor in the Generation of Nitrogen Oxides by Murine Macrophages," The Journal of Biological Chemistry, Vol. 264, No. 34, pp. 20496-20501, (1989).	
		GIOVANELLI, J. et al., "Tetrahydrobiopterin, a Cofactor for Rat Cerebellar Nitric Does Not Function as a Reactant in the Oxygenation of Arg," Proc. Natl. Acad. Sci., Vol. 88, No. 16, p. 7091, (1991). (Abstract Only)	
		MÜLSCH, A. et al., "Nitric Oxide Synthase in Native and Cultured Endothelial Cells: Calcium/Calmodulin and Tetrahydrobiopterin Are Cofactors," Journal of Cardiovascular Pharmacology, Vol. 17, Suppl. 3, pp. S52-S56, (1991).	
		SAKAI, N. et al., "Tetrahydrobiopterin is Required for Cytokine-Induced Nitric Oxide Production in a Murine Macrophage Cell Line (RAW 264)," Mol. Pharmacol., Vol. 43, Issue 1, p. 6, (1993). (Abstract Only)	
		KLATT, P. et al., "Stimulation of Human Nitric Oxide Synthase by Tetrahydrobiopterin and Selective Binding of the Cofactor," FEBS Letters, Vol. 305, No. 2, pp. 160-162, (1992).	
		WERNER-FELMAYER, G. et al., "Ca <sup>2+</sup> /Calmodulin-Dependent Nitric Oxide Synthase Activity in the Human Cervix Carcinoma Cell Line ME-180," Biochem. J., Vol. 289, pp. 357-361, (1993).	
		HEVEL, J. M. et al., "Macrophage Nitric Oxide Synthase: Relationship Between Enzyme-Bound Tetrahydrobiopterin and Synthase Activity," Biochemistry, Vol. 31, pp. 7160-7165, (1992).	
		PFEIFFER, S. et al., "Allosteric Modulation of Rat Brain Nitric Oxide Synthase by the Pterin-Site Enzyme Inhibitor 4-Aminotetrahydrobiopterin," Biochem. J., Vol. 328, pp. 349-352, (1997).	
		WERNER, E. R. et al., "Identification of the 4-Amino Analogue of Tetrahydrobiopterin as a Dihydropteridine Reductase Inhibitor and a Potent Pteridine Antagonist of Rat Neuronal Nitric Oxide Synthase," Biochem. J., Vol. 320, pp. 193-196, (1996).	

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Sheet	2	of	2	Attorney Docket Number	05281.0018

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		SCHIRCKS, V. B. et al., "A New, Regiospecific Synthesis of L-Biopterine," Helvetica Chimica Acta, Vol. 60, No. 1, pp. 211-214, (1977).		Abstract
		FUTTERMAN, S., "Enzymatic Reduction of Folic Acid and Dihydrofolic Acid to Tetrahydro-Folic Acid," J. Biol. Chem., Vol. 228, pp. 1031-1038, (1957). (Abstract Only)		
		FUKUSHIMA, T. et al., "Nuclear Magnetic Resonance Studies of Some Biologically Active Dihydropterins," Vol. 128, Issue 1, 1 page, (1968).		
		PFLEIDERER, W. et al., "A Simple Synthetic Approach to 8-Substituted 5,6,7,8-Tetrahydro- and 7,8-Dihydropterins," Chem. Ber., Vol. 104, pp. 2293-2312, (1971).		YES
		ANDREWS, K. J. M. et al., "A New Synthesis of Biopterin and L-Neopterin," Chemical Communications, pp. 120-121, (1968).		
		HANAYA, T. et al., "Pteridines CV Selective N(3)-and O <sup>4</sup> -Alkylation of L-Biopterin: A Convenient Synthesis of 3-and O <sup>4</sup> -Methyl-L-Biopterin and the Versatile N <sup>2</sup> -(N,N-Dimethylaminomethylene)-N(3)-p-Nitrophenethyl-Protected L-Biopterin," Pteridines, Vol. 6, No. 1, pp. 1-7, (1995).		
		MATTER, H. et al., "Structural Requirements for Inhibition of the Neuronal Nitric Oxide synthase (NOS-I): 3D-QSAR Analysis of 4-Oxo-and 4-Amino-Pteridine-Based Inhibitors," J. Med. Chem., Vol. 45, No. 14, pp. 2923-2941, (2002).		
		KNIPP, M. et al., "A Colorimetric 96-Well Microtiter Plate Assay for the Determination of Enzymatically Formed Citrulline," Anal Biochem., Vol. 286, No. 2, pp. 257-64, (2000). (Abstract Only)		
		TAYLOR, E. C. et al., "Pteridines. XXIX. An Unequivocal Route to 2,4-Diamino-6-Substituted Pteridines," Journal of the American Chemical Society, Vol. 95, No. 19, pp. 6413-6418, (1973).		
		KWEE, S. et al., "Electrochemistry of Some Substituted Pteridines," Biochimica et Biophysica Acta, Vol. 297, No. 2, p 285-296, (1973). (Abstract Only)		
		ZIMMERMAN, M. et al., "Inhibitors of Folate Biosynthesis. 1. Inhibition of Dihydroneopterin Aldolase by Pteridine Derivatives," Journal of Medicinal Chemistry, Vol. 20, No. 9, pp. 1213-1215, (1977). (Abstract Only)		
		AL-HASSAN et al., "2,4-Diamino-7,8-dihydro-6,7,7-trimethylpteridine," J. Chem. Soc. Perkin Trans 1, pp. 2145-2150, (1985).		
		AL-HASSAN et al., "2,4-Diamino-8-benzyl-7,8-dihydro-6,7,7-trimethylpteridine," J. Chem. Soc. Perkin Trans 1, pp. 2145-2150, (1985).		
		Chaykovsky et al., "6-methyl-7,8-dihydro-pteridine-2,4-diamine," J. Org. Chem., Vol. 40, p. 145, (1975).		

Examiner Signature		Date Considered	
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EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.